

A32040 PCT USA-A - 072876.0102

PATENT

*G1  
F1  
cancer*

or a cell cycle affecting gene operably linked to

(ii) a cell type specific promoter,

in which the conditional oncogene, transforming gene or immortalizing gene or the cell cycle affecting gene is a SV40tsA58 gene

and in which the cell type specific promoter is a human NF-L gene promoter.

*F2*

7. (Fourth Amendment) A neuronal cell line obtained from a transgenic rat, the cells of which comprise:

(i) a conditional oncogene, transforming gene or immortalizing gene or a cell cycle affecting gene operably linked to

(ii) a cell type specific promoter

in which the conditional oncogene, transforming gene, immortalizing gene or the cell cycle affecting gene is a C Erb  $\beta 2$  gene or a TGF  $\alpha$  gene

and in which the cell type specific promoter is a human NF-L gene promoter.

*F3*

9. (Amended) A cell line as claimed in claim 1 having the ECACC Accession number 96092454.

*F4  
G1*

13. (Thrice Amended) A method of producing a transgenic rat, comprising:

(i) causing a female rat to super-ovulate by supplying her with a

A32040 PCT USA-A - 072876.0102

PATENT

regular supply of Follicle Stimulating Hormone (FSH) prior to mating;

(ii) mating or artificially inseminating the female rat;

(iii) obtaining the resulting embryo from the female rat; and

(iv) incorporating

(i) a conditional oncogene, transforming gene or

immortalizing gene or a cell cycle affecting gene operably linked to

(ii) a cell specific promoter into the genome of the rat embryo

in which the conditional oncogene, transforming gene or immortalizing gene or the cell cycle affecting gene is a SV40tsA58 gene, C Erb  $\beta$ 2 gene or a TGF  $\alpha$  gene

and in which the cell type specific promoter is a human NF-L gene

promoter.

17. (Thrice Amended) A transgenic rat whose germ cells and somatic cells contain

(i) a conditional oncogene, transforming gene or immortalizing gene or a cell cycle affecting gene operably linked to

(ii) a cell type specific promoter as a result of chromosomal incorporation into the rat genome or into the genome of an ancestor of said rat

in which the conditional oncogene, transforming gene or immortalizing gene or the cell cycle affecting gene is a SV40tsA58 gene

and in which the cell type specific promoter is a human NF-L gene

A32040 PCT USA-A - 072876.0102

PATENT

~~promoter.~~

18. (Thrice Amended) A transgenic rat whose germ cells and somatic cells contain

(i) a conditional oncogene, transforming gene or immortalising gene or a cell cycle affecting gene operably linked to

(ii) a cell type specific promoter as a result of chromosomal incorporation into the rat genome or into the genome of an ancestor of said rat,

wherein the conditional oncogene, transforming gene, immortalising gene, or the cell cycle affecting gene is a C Erb  $\beta$  2 gene or a TGF  $\alpha$  gene,

and wherein the cell type specific promoter is a human NF-L gene promoter.

25. (Twice Amended) A method of generating a cell line from a transgenic rat comprising a conditional oncogene, transforming gene or immortalizing gene or a cell cycle affecting gene operably linked to a cell specific promoter wherein the cell type specific promoter is a human NF-L gene promoter, the method comprising:

(i) maintaining the rat at restrictive conditions such that the conditional oncogene, transforming gene or immortalizing gene or the cell cycle affecting gene is a SV40tsA58 gene, a C Erb  $\beta$  2 gene, or a TGF  $\alpha$  gene and is expressed in vivo, only in a tissue of interest and in an inactive form such that the cells thereof grow normally;